

U.S.S.N. 10/099,830

U.S. Pat. No. 6,559,626

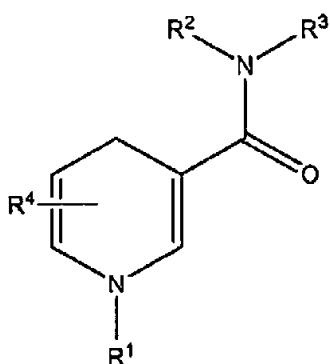
AMENDMENT AND RESPONSE TO OFFICE ACTION

### Amendment

### In the Claims

Claims 1-33 (Canceled).

34. (Twice Amended) A therapeutic system comprising a prodrug which is converted to a substantially cytotoxic drug by the action of  $\text{NO}_2$  and a compound of formula I.



wherein R<sup>1</sup> is selected from the group consisting of substituted alkyl, including substitution by CONH<sub>2</sub>, OH, halogen, CN, and COOH; aryl; substituted aryl; CONR<sup>a</sup>R<sup>b</sup>, where R<sup>a</sup> and R<sup>b</sup> are independently H, alkyl, or substituted alkyl, and R<sup>2</sup> and R<sup>3</sup> are independently H, alkyl, or substituted alkyl and R<sup>4</sup> is any of H, alkyl, substituted alkyl, halogen, CN, COOH, CONH<sub>2</sub>, or OH, wherein the compound can pass reducing equivalents to NQO<sub>2</sub>, in a form for administration to a patient in need thereof.

Claims 35-40 (Canceled)

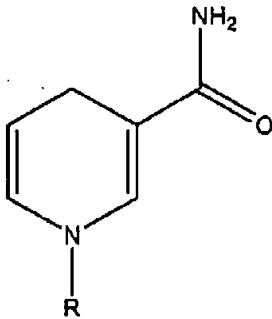
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**AMENDMENT AND RESPONSE TO OFFICE ACTION**

41. (Previously Presented) The system of claim 34, wherein the compound is 1-(carboxamidomethyl)-dihydronicotinamide.

42. (Currently Amended) The system of claim 34, wherein the compound has formula II



wherein R is a substituted alkyl, comprising one or more groups selected from the group consisting of ~~CNH<sub>2</sub> CONH<sub>2</sub>~~, OH, halogen, CN, and COOH.

43. (Previously Presented) The system of claim 34, wherein the alkyl group is a C<sub>1</sub> to C<sub>6</sub> alkyl.

44. (Currently Amended) The system of claim 34 42, wherein R is selected from the group consisting of -CH<sub>2</sub>CONH<sub>2</sub>, -CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>SO<sub>3</sub><sup>-</sup>, -CH<sub>2</sub>CH<sub>2</sub>OH, -CH<sub>2</sub>CH<sub>2</sub>CH<sub>2</sub>OH, and -CH<sub>2</sub>CH<sub>2</sub>COOH.

45. (Canceled) The system of claim 34 wherein the analogue of NRH is 1-(carboxamidomethyl)-dihydronicotinamide.

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46. (New) The therapeutic system of claim 34, wherein the prodrug is CB 1954 or an analogue thereof

47. (New) The therapeutic system of claim 34, wherein the prodrug is CB 1954.

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